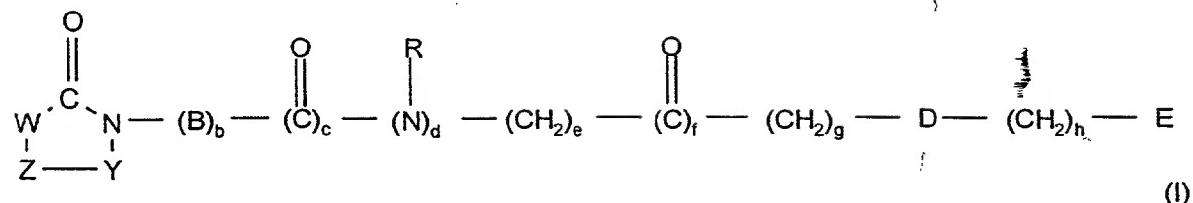


What is claimed is:

1. A compound of the formula I



in which

- W is $R^1\text{-A-C}(R^{13})$ or $R^1\text{-A-CH=}$ C;
- Y is a carbonyl, thiocarbonyl or methylene group;
- Z is $N(R^0)$, oxygen, sulfur or a methylene group;
- A is a bivalent radical from the group consisting of ($C_1\text{-}C_6$)-alkylene, ($C_3\text{-}C_{12}$)-cycloalkylene, ($C_1\text{-}C_6$)-alkylene-($C_3\text{-}C_{12}$)-cycloalkyl, phenylene, phenylene-($C_1\text{-}C_6$)-alkyl, ($C_1\text{-}C_6$)-alkylenephenoxy, ($C_1\text{-}C_6$)-alkylenephenoxy-($C_1\text{-}C_6$)-alkyl, phenylene-($C_2\text{-}C_6$)-alkenyl or a bivalent radical of a 5- or 6-membered saturated or unsaturated ring which can contain 1 or 2 nitrogen atoms and can be mono- or disubstituted by ($C_1\text{-}C_6$)-alkyl or doubly bonded oxygen or sulfur, or is a direct bond;
- B is a bivalent radical from the group consisting of ($C_1\text{-}C_6$)-alkylene, ($C_2\text{-}C_6$)-alkenylene, phenylene, phenylene-($C_1\text{-}C_3$)-alkyl, ($C_1\text{-}C_3$)-alkylenephenoxy, where the bivalent ($C_1\text{-}C_6$)-alkylene radical can be unsubstituted or substituted by a radical from the group consisting of ($C_1\text{-}C_8$)-alkyl, ($C_2\text{-}C_8$)-alkenyl, ($C_2\text{-}C_8$)-alkynyl, ($C_3\text{-}C_{10}$)-cycloalkyl, ($C_3\text{-}C_{10}$)-cycloalkyl-($C_1\text{-}C_6$)-alkyl, optionally substituted ($C_6\text{-}C_{14}$)-aryl, ($C_6\text{-}C_{14}$)-aryloxy, ($C_1\text{-}C_6$)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl and heteroaryl-($C_1\text{-}C_6$)-alkyl optionally substituted in the heteroaryl radical;
- D is $C(R^2)(R^3)$, $N(R^3)$ or $CH=C(R^3)$;
- E is tetrazolyl, $(R^8O)_2P(O)$, $HOS(O)_2$, $R^9NHS(O)_2$ or $R^{10}CO$;
- R is hydrogen, ($C_1\text{-}C_8$)-alkyl, ($C_3\text{-}C_{12}$)-cycloalkyl, ($C_3\text{-}C_{12}$)-cycloalkyl-($C_1\text{-}C_8$)-alkyl, optionally substituted ($C_6\text{-}C_{14}$)-aryl, ($C_6\text{-}C_{14}$)-aryl-($C_1\text{-}C_8$)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl and heteroaryl-($C_1\text{-}C_8$)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be mono- or polysubstituted by fluorine;
- R^0 is hydrogen, ($C_1\text{-}C_8$)-alkyl, ($C_3\text{-}C_{12}$)-cycloalkyl, ($C_3\text{-}C_{12}$)-cycloalkyl-($C_1\text{-}C_8$)-alkyl, ($C_6\text{-}C_{12}$)-bicycloalkyl, ($C_6\text{-}C_{12}$)-bicycloalkyl-($C_1\text{-}C_8$)-alkyl, ($C_6\text{-}C_{12}$)-tricycloalkyl, ($C_6\text{-}C_{12}$)-tricycloalkyl-($C_1\text{-}C_8$)-alkyl, optionally substituted ($C_6\text{-}C_{14}$)-aryl, ($C_6\text{-}C_{14}$)-aryl-($C_1\text{-}C_8$)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-($C_1\text{-}C_8$)-alkyl optionally substituted in the heteroaryl radical, CHO, ($C_1\text{-}C_8$)-alkyl-CO, ($C_3\text{-}C_{12}$)-cycloalkyl-CO, ($C_3\text{-}C_{12}$)-cycloalkyl-($C_1\text{-}C_8$)-alkyl-CO, ($C_6\text{-}C_{12}$)-bicycloalkyl-CO, ($C_6\text{-}C_{12}$)-bicycloalkyl-($C_1\text{-}C_8$)-alkyl-CO, ($C_6\text{-}C_{12}$)-tricycloalkyl-CO,

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(C₆-C₁₂)-tricycloalkyl-(C₁-C₈)-alkyl-CO, optionally substituted (C₆-C₁₄)-aryl-CO, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl-CO optionally substituted in the aryl radical, optionally substituted heteroaryl-CO, heteroaryl-(C₁-C₈)-alkyl-CO optionally substituted in the heteroaryl radical, (C₁-C₈)-alkyl-S(O)_n, (C₃-C₁₂)-cycloalkyl-S(O)_n, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl-S(O)_n, (C₆-C₁₂)-bicycloalkyl-S(O)_n, (C₆-C₁₂)-bicycloalkyl-(C₁-C₈)-alkyl-S(O)_n, (C₆-C₁₂)-tricycloalkyl-S(O)_n, (C₆-C₁₂)-tricycloalkyl-(C₁-C₈)-alkyl-S(O)_n, optionally substituted (C₆-C₁₄)-aryl-S(O)_n, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl-S(O)_n, optionally substituted in the aryl radical, optionally substituted heteroaryl-S(O)_n or heteroaryl-(C₁-C₈)-alkyl-S(O)_n optionally substituted in the heteroaryl radical, where n is 1 or 2;

10 R¹

is one of the radicals -S-R²¹, -S-S-R²¹, -S(O)-R²², -S(O)-R²², -S-OR²¹, -S(O)-OR²¹, -S(O)-OR²¹, -S-N(R²¹)-R²⁸, -S(O)-N(R²¹)-R²⁸, -S(O)-N(R²¹)-R²⁸, -S-C(O)-R²¹, -S-C(O)-OR²², -S-C(S)-SR²², -S-C(O)-N(R²¹)-R²⁸, -S-C(S)-N(R²¹)-R²⁸, -O-C(O)-R²¹, -O-C(S)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸, -O-S(O)-OR²¹, -O-S(O)-OR²¹, -O-S(O)-R²², -O-S(O)-R²², -O-P(O)(OR²¹)₂, -O-P(O)(OR²¹)-N(R²¹)-R²⁸, -O-P(O)(N(R²¹)-R²⁸)₂, -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(O)-SR²², -N(R²⁸)-C(S)-OR²², -N(R²⁸)-C(S)-SR²², -N(R²⁸)-C(S)-R²¹, -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)-OR²¹, -N(R²⁸)-S(O)-OR²¹, -N(R²⁸)-S(O)-N(R²¹)-R²⁸, -N(R²⁸)-P(O)(OR²¹)₂, -N(R²⁸)-P(O)(OR²¹)-N(R²¹)-R²⁸, -N(R²⁸)-P(O)(N(R²¹)-R²⁸)₂, -N(R²⁸)-P(O)(R²²)-OR²¹, -N(R²⁸)-P(O)(R²²)-OR²¹, -P(O)(OR²¹)₂, -P(O)(OR²¹)-N(R²¹)-R²⁸, -P(O)(N(R²¹)-R²⁸)₂, -P(O)(R²²)-OR²¹, -P(O)(R²²)-N(R²¹)-R²⁸, -P(O)(R²²)₂, -C(S)-R²¹, -C(S)-SR²¹, -C(S)-N(R²¹)-R²⁸,

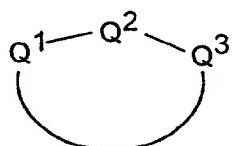
15

cyano, halogen, nitro or methylenedioxy or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula

20

25

30



in which

Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂;

Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-),

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-) representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A,

the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

- 5 R² is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl;
- 10 R³ is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, (C₃-C₈)-cycloalkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₂-C₈)-alkenylcarbonyl, (C₂-C₈)-alkynylcarbonyl, pyridyl, R¹¹NH, R⁴CO, COOR⁴, CON(CH₃)R⁴, CONHR⁴, CSNHR⁴, COOR¹⁵, CON(CH₃)R¹⁵ or CONHR¹⁵;
- 15 R⁴ is hydrogen or (C₁-C₂₈)-alkyl which can optionally be mono- or polysubstituted by identical or different radicals from the group consisting of hydroxyl, hydroxycarbonyl, aminocarbonyl, mono- or di-((C₁-C₁₈)-alkyl)aminocarbonyl, amino-(C₂-C₁₈)-alkylaminocarbonyl, amino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₂-C₁₈)-alkylaminocarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxycarbonyl which can also be substituted in the aryl radical, amino, mercapto, (C₁-C₁₈)-alkoxy, (C₁-C₁₈)-alkoxycarbonyl, Het-CO, optionally substituted (C₃-C₈)-cycloalkyl, HOS(O)₂-(C₁-C₃)-alkyl, R⁹NHS(O)₂-(C₁-C₃)-alkyl, (R⁸O)₂P(O)-(C₁-C₃)-alkyl, tetrazolyl-(C₁-C₃)-alkyl, halogen, nitro, trifluoromethyl or the radical R⁵;
- 20 R⁵ is optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, a mono- or bicyclic 5- to 12-membered heterocyclic ring which can be aromatic, partially hydrogenated or completely hydrogenated and which can contain one, two or three identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur, a radical R⁶ or a radical R⁶CO-, where the aryl radical and, independently thereof, the heterocyclic radical can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₁₈)-alkyl, (C₁-C₁₈)-alkoxy, halogen, nitro, amino and trifluoromethyl;
- 25 R⁶ is R⁷R⁸N, R⁷O or R⁷S or an amino acid side chain, a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-, and their esters and amides, where hydrogen or hydroxymethyl can optionally stand in place of free functional groups and/or where free functional groups can be protected by protective groups customary in peptide chemistry;
- 30 R⁷ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-;
- 35 R⁸ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-;
- 40 R⁹ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-;

	R^8	is hydrogen, $(C_1\text{-}C_{18})\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$ or $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$ which can also be substituted in the aryl radical;
5	R^9	is hydrogen, aminocarbonyl, $(C_1\text{-}C_{18})\text{-alkylaminocarbonyl}$, $(C_3\text{-}C_8)\text{-cycloalkylaminocarbonyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-arylaminocarbonyl}$, $(C_1\text{-}C_{18})\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$ or $(C_3\text{-}C_8)\text{-cycloalkyl}$;
10	R^{10}	is hydroxyl, $(C_1\text{-}C_{18})\text{-alkoxy}$, $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkoxy}$ which can also be substituted in the aryl radical, optionally substituted $(C_6\text{-}C_{14})\text{-aryloxy}$, amino or mono- or di- $((C_1\text{-}C_{18})\text{-alkyl})\text{amino}$;
15	R^{11}	is hydrogen, R^{12a} , $R^{12a}\text{-CO}$, $H\text{-CO}$, $R^{12a}\text{-O-CO}$, $R^{12b}\text{-CO}$, $R^{12b}\text{-CS}$, $R^{12a}\text{-S(O)}_2$ or $R^{12b}\text{-S(O)}_2$;
20	R^{12a}	is $(C_1\text{-}C_{18})\text{-alkyl}$, $(C_2\text{-}C_8)\text{-alkenyl}$, $(C_2\text{-}C_8)\text{-alkynyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$, $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl- $(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the heteroaryl radical or the radical R^{15} ;
25	R^{12b}	is amino, di- $((C_1\text{-}C_{18})\text{-alkyl})\text{amino}$ or $R^{12a}\text{-NH}$;
30	R^{13}	is hydrogen, $(C_1\text{-}C_6)\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$, $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the aryl radical, $(C_3\text{-}C_8)\text{-cycloalkyl}$ or $(C_3\text{-}C_8)\text{-cycloalkyl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$;
35	R^{15}	is $R^{16}\text{-}(C_1\text{-}C_6)\text{-alkyl}$ or R^{16} ;
40	R^{16}	is a 6- to 24-membered bicyclic or tricyclic radical which is saturated or partially unsaturated and which can also contain one, two, three or four identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur and which can also be substituted by one or more identical or different substituents from the group consisting of $(C_1\text{-}C_4)\text{-alkyl}$ and oxo;
	R^{21}	is hydrogen, $(C_1\text{-}C_8)\text{-alkyl}$, hydroxy- $(C_1\text{-}C_8)\text{-alkyl}$, $(C_2\text{-}C_8)\text{-alkenyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$, $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl- $(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the heteroaryl radical, where alkyl radicals can also be monosubstituted or polysubstituted by fluorine and the radicals R^{21} can be identical or different if they occur two or more times;
	R^{22}	is $(C_1\text{-}C_8)\text{-alkyl}$, hydroxy- $(C_1\text{-}C_8)\text{-alkyl}$, $(C_2\text{-}C_8)\text{-alkenyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}$, $(C_3\text{-}C_{12})\text{-cycloalkyl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$, optionally substituted $(C_6\text{-}C_{14})\text{-aryl}$, $(C_6\text{-}C_{14})\text{-aryl}\text{-}(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl- $(C_1\text{-}C_8)\text{-alkyl}$ optionally substituted in the heteroaryl radical, where alkyl radicals can be monosubstituted or polysubstituted by fluorine and the radicals R^{22} can be identical or different if they occur two or more times;
	R^{28}	is one of the radicals $R^{21}\text{-}$, $R^{21}\text{N}(R^{21})\text{-}$, $R^{21}\text{C(O)}\text{-}$, $R^{22}\text{O-C(O)}\text{-}$, $R^{21}\text{N}(R^{21})\text{-C(O)}\text{-}$ or $R^{21}\text{N}(R^{21})\text{-C(=N(R^{21}))}\text{-}$;
	R^{29}	is one of the radicals $R^{22}\text{-}$, $R^{21}\text{N}(R^{21})\text{-}$, $R^{21}\text{C(O)}\text{-}$, $R^{22}\text{O-C(O)}\text{-}$, $R^{21}\text{N}(R^{21})\text{-C(O)}\text{-}$ or $R^{21}\text{N}(R^{21})\text{-C(=N(R^{21}))}\text{-}$;
	Het	is the radical of a 5- to 10-membered, monocyclic or polycyclic heterocycle bonded via

a nitrogen atom, which can be aromatic or partially unsaturated or saturated and which can contain one, two, three or four identical or different additional ring heteroatoms from the group consisting of oxygen, nitrogen and sulfur and which can be optionally substituted on carbon atoms and on additional ring nitrogen atoms, where there can be identical or different radicals R^h , R^hCO or R^hO-CO as substituents on additional ring nitrogen atoms and R^h is hydrogen, (C_1-C_8) -alkyl, (C_3-C_8) -cycloalkyl, (C_3-C_8) -cycloalkyl- (C_1-C_8) -alkyl, optionally substituted (C_6-C_{14}) -aryl or (C_6-C_{14}) -aryl- (C_1-C_8) -alkyl optionally substituted in the aryl radical;

b, c, d and f independently of one another are 0 or 1, but cannot all simultaneously be 0;

e, g and h independently of one another are 0, 1, 2, 3, 4, 5 or 6; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts;

where, however, if simultaneously W is 4-cyanophenyl- $C(R^{13})$, Y is a carbonyl group, Z is NR^{0a} , B is an unsubstituted methylene group, R is R^a , b, c and d are 1 and e, f and g are 0, then D cannot be $C(R^{2a})(R^{3a})$, where

R^{0a} , R^a and R^{2a} independently of one another are hydrogen, (C_1-C_8) -alkyl, optionally substituted (C_6-C_{14}) -aryl, (C_6-C_{14}) -aryl- (C_1-C_8) -alkyl optionally substituted in the aryl radical or (C_3-C_8) -cycloalkyl and

R^{3a} is hydrogen, (C_1-C_8) -alkyl, optionally substituted (C_6-C_{14}) -aryl, (C_6-C_{14}) -aryl- (C_1-C_8) -alkyl optionally substituted in the aryl radical, (C_3-C_8) -cycloalkyl or 2-, 3- or 4-pyridyl.

2. A compound of the formula I as claimed in claim 1, in which

W is $R^1-A-C(R^{13})$ or $R^1-A-CH=C$;

Y is a carbonyl, thiocarbonyl or methylene group;

Z is $N(R^0)$, oxygen, sulfur or a methylene group;

A is a bivalent radical from the group consisting of (C_1-C_6) -alkylene, (C_3-C_{12}) -cycloalkylene, (C_1-C_6) -alkylene- (C_3-C_{12}) -cycloalkyl, phenylene, phenylene- (C_1-C_6) -alkyl, (C_1-C_6) -alkylenephenoxy, (C_1-C_6) -alkylenephenoxy- (C_1-C_6) -alkyl, phenylene- (C_2-C_6) -alkenyl or a bivalent radical of a 5- or 6-membered saturated or unsaturated ring which can contain 1 or 2 nitrogen atoms and can be mono- or disubstituted by (C_1-C_6) -alkyl or doubly bonded oxygen or sulfur, or is a direct bond;

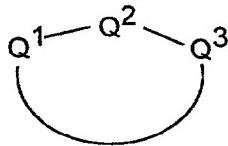
B is a bivalent radical from the group consisting of (C_1-C_6) -alkylene, (C_2-C_6) -alkenylene, phenylene, phenylene- (C_1-C_3) -alkyl, (C_1-C_3) -alkylenephenoxy;

D is $C(R^2)(R^3)$, $N(R^3)$ or $CH=C(R^3)$;

E is tetrazolyl, $(R^8O)_2P(O)$, $HOS(O)_2$, $R^9NHS(O)_2$ or $R^{10}CO$;

R and R^0 independently of one another are hydrogen, (C_1-C_8) -alkyl, (C_3-C_{12}) -cycloalkyl, (C_3-C_{12}) -cycloalkyl- (C_1-C_8) -alkyl, optionally substituted (C_6-C_{14}) -aryl, (C_6-C_{14}) -aryl- (C_1-C_8) -alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl- (C_1-C_8) -alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be mono- or polysubstituted by fluorine;

R¹ is one of the radicals -S-R²¹, -S-S-R²¹, -S(O)-R²², -S(O)₂-R²², -S-OR²¹, -S(O)-OR²¹, -S(O)₂-OR²¹, -S-N(R²¹)-R²⁸, -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸, -S-C(O)-R²¹, -S-C(O)-OR²², -S-C(S)-SR²², -S-C(O)-N(R²¹)-R²⁸, -S-C(S)-N(R²¹)-R²⁸, -O-C(O)-R²¹, -O-C(S)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸,
5 -O-C(S)-N(R²¹)-R²⁸, -O-S(O)₂-OR²¹, -O-S(O)-OR²¹, -O-S(O)₂-N(R²¹)-R²⁸,
-O-S(O)-N(R²¹)-R²⁸, -O-S(O)₂-R²², -O-S(O)-R²²,
-O-P(O)(OR²¹)₂, -O-P(O)(OR²¹)-N(R²¹)-R²⁸, -O-P(O)(N(R²¹)-R²⁸)₂,
-N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(O)-SR²², -N(R²⁸)-C(S)-OR²²,
-N(R²⁸)-C(S)-SR²², -N(R²⁸)-C(S)-R²¹,
10 -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸,
-N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹, -N(R²⁸)-S(O)₂-N(R²¹)-R²⁸,
15 -N(R²⁸)-P(O)(OR²¹)₂, -N(R²⁸)-P(O)(OR²¹)-N(R²¹)-R²⁸,
-N(R²⁸)-P(O)(N(R²¹)-R²⁸)₂, -N(R²⁸)-P(O)(R²²)-OR²¹,
-N(R²⁸)-P(O)(R²²)-N(R²¹)-R²⁸, -N(R²⁸)-P(O)(R²²)₂,
20 -P(O)(OR²¹)₂, -P(O)(OR²¹)-N(R²¹)-R²⁸, -P(O)(N(R²¹)-R²⁸)₂, -P(O)(R²²)-OR²¹,
-P(O)(R²²)-N(R²¹)-R²⁸, -P(O)(R²²)₂,
-C(S)-R²¹, -C(S)-SR²¹, -C(S)-N(R²¹)-R²⁸,
25 cyano, halogen, nitro or methylenedioxy or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



in which

Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂-;

Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-)- representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

R² is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl;

R³ is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, (C₃-C₈)-cycloalkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₂-C₈)-alkenylcarbonyl, (C₂-C₈)-alkynylcarbonyl, pyridyl, R¹¹NH, R⁴CO, COOR⁴, CON(CH₃)R⁴, CONHR⁴, CSNHR⁴, COOR¹⁵, CON(CH₃)R¹⁵ or CONHR¹⁵;

R⁴ is hydrogen or (C₁-C₂₈)-alkyl which can optionally be mono- or polysubstituted by

identical or different radicals from the group consisting of hydroxyl, hydroxycarbonyl, aminocarbonyl, mono- or di-((C₁-C₁₈)-alkyl)aminocarbonyl, amino-(C₂-C₁₈)-alkylaminocarbonyl, amino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl,
 5 (C₁-C₁₈)-alkylcarbonylamino-(C₂-C₁₈)-alkylaminocarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxycarbonyl which can also be substituted in the aryl radical, amino, mercapto, (C₁-C₁₈)-alkoxy, (C₁-C₁₈)-alkoxycarbonyl, optionally substituted (C₃-C₈)-cycloalkyl, HOS(O)₂-(C₁-C₃)-alkyl, R⁹NHS(O)₂-(C₁-C₃)-alkyl, (R⁸O)₂P(O)-(C₁-C₃)-alkyl, tetrazolyl-(C₁-C₃)-alkyl, halogen, nitro, trifluoromethyl or the radical R⁵;

10 R⁵ is optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, a mono- or bicyclic 5- to 12-membered heterocyclic ring which can be aromatic, partially hydrogenated or completely hydrogenated and which can contain one, two or three identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur, a radical R⁶ or a radical R⁶CO-, where the aryl radical and, independently thereof, the heterocyclic radical can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₁₈)-alkyl, (C₁-C₁₈)-alkoxy, halogen, nitro, amino or trifluoromethyl;

15 R⁶ is R⁷R⁸N, R⁷O or R⁷S or an amino acid side chain, a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or in which the peptide bond can be reduced to -NH-CH₂-, and their esters and amides, where hydrogen or hydroxymethyl can optionally stand in place of free functional groups and/or where free functional groups can be protected by protective groups customary in peptide chemistry;

20 R⁷ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted, by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or in which the peptide bond can be reduced to -NH-CH₂-;

25 R⁸ is hydrogen, (C₁-C₁₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl which can also be substituted in the aryl radical;

30 R⁹ is hydrogen, aminocarbonyl, (C₁-C₁₈)-alkylaminocarbonyl, (C₃-C₈)-cycloalkylaminocarbonyl, optionally substituted (C₆-C₁₄)-arylamino carbonyl, (C₁-C₁₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₃-C₈)-cycloalkyl;

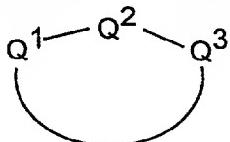
35 R¹⁰ is hydroxyl, (C₁-C₁₈)-alkoxy, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxy which can also be substituted in the aryl radical, optionally substituted (C₆-C₁₄)-aryloxy, amino or mono- or di-((C₁-C₁₈)-alkyl)amino;

40 R¹¹ is hydrogen, (C₁-C₁₈)-alkyl, R¹²CO, optionally substituted (C₆-C₁₄)-aryl-S(O)₂, (C₁-C₁₈)-

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- alkyl-S(O)₂, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, R⁹NHS(O)₂ or the radical R¹⁵;
- R¹² is hydrogen, (C₁-C₁₈)-alkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, optionally substituted (C₆-C₁₄)-aryl, (C₁-C₁₈)-alkoxy, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxy which can also be substituted in the aryl radical, optionally substituted (C₆-C₁₄)-aryloxy, amino or mono- or di-((C₁-C₁₈)-alkyl)amino, the radical R¹⁵ or the radical R¹⁵-O-;
- R¹³ is hydrogen, (C₁-C₆)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl;
- R¹⁵ is R¹⁶-(C₁-C₆)-alkyl or R¹⁶;
- R¹⁶ is a 6- to 24-membered bicyclic or tricyclic radical which is saturated or partially unsaturated and which can also contain one to four identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur and which can also be substituted by one or more identical or different substituents from the group consisting of (C₁-C₄)-alkyl and oxo;
- R²¹ is hydrogen, (C₁-C₈)-alkyl, hydroxy-(C₁-C₈)-alkyl, (C₂-C₈)-alkenyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be monosubstituted or polysubstituted by fluorine and the radicals R²¹ can be identical or different if they occur two or more times;
- R²² is (C₁-C₈)-alkyl, hydroxy-(C₁-C₈)-alkyl, (C₂-C₈)-alkenyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be monosubstituted or polysubstituted by fluorine and the radicals R²² can be identical or different if they occur two or more times;
- R²⁸ is one of the radicals R²¹-, R²¹N(R²¹)-, R²¹C(O)-, R²²O-C(O)-, R²¹N(R²¹)-C(O)- or R²¹N(R²¹)-C(=N(R²¹))-;
- R²⁹ is one of the radicals R²²-, R²¹N(R²¹)-, R²¹C(O)-, R²²O-C(O)-, R²¹N(R²¹)-C(O)- or R²¹N(R²¹)-C(=N(R²¹))-;
- b, c, d and f independently of one another are 0 or 1, but cannot all simultaneously be 0; e, g and h independently of one another are 0, 1, 2, 3, 4, 5 or 6; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.
3. A compound of the formula I as claimed in claim 1, in which R¹ is one of the radicals -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸, -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸, -O-S(O)₂-N(R²¹)-R²⁸, -O-S(O)-N(R²¹)-R²⁸, -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(S)-R²¹, -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸, -N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹,

-N(R²⁸)-S(O)₂-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-N(R²¹)-R²⁸,

-C(S)-R²¹, -C(S)-N(R²¹)-R²⁸ or cyano or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



5 in which

Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂;

Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-),

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-) representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

15 4. A compound of the formula I as claimed in claim 1, in which R⁰ is (C₁-C₈)-alkyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

25 5. A compound of the formula I as claimed in claim 4, wherein R⁰ is selected from the group consisting of biphenylylmethyl, naphthylmethyl, and benzyl, each of which is unsubstituted or monosubstituted or polysubstituted in the aryl radical; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

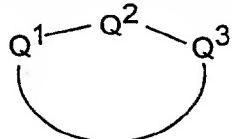
30 6. A compound of the formula I as claimed in claim 1, in which simultaneously W is R¹-A-CH=C and therein A is a phenylene radical or a methylenephenoxy radical, or W is R¹-A-C(R¹³) and therein A is a bivalent radical from the group consisting of methylene, ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene, phenylenemethyl, methylenephenoxy, methylenephenoxyethyl; B is a bivalent radical from the group consisting of methylene, ethylene, trimethylene, tetramethylene, vinylene, phenylene, or substituted methylene or ethylene; E is R¹⁰CO; R is hydrogen, (C₁-C₆)-alkyl or benzyl; R⁰ is (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-ary-

(C₁-C₈)-alkyl optionally substituted in the aryl radical;

R¹ is one of the radicals -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸,
 -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸,
 -O-S(O)₂-N(R²¹)-R²⁸, -O-S(O)-N(R²¹)-R²⁸,

5 -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(S)-R²¹,
 -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸,
 -N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹,
 -N(R²⁸)-S(O)₂-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-N(R²¹)-R²⁸,

10 -C(S)-R²¹, -C(S)-N(R²¹)-R²⁸ or cyano or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



in which

Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂;

15 Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-) representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

20 R² is hydrogen or (C₁-C₈)-alkyl.

R³ is (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl,
 (C₃-C₈)-cycloalkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, pyridyl, R¹¹NH, R⁴CO, COOR⁴,
 CONHR⁴, CSNHR⁴, COOR¹⁵ and CONHR¹⁵;

and e, g and h independently of one another are the numbers 0, 1, 2 or 3;

25 in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

7. A compound of the formula I as claimed in claim 1, in which W is R¹-A-C(R¹³) and R¹³ is (C₁-C₆)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

8. A compound of the formula I as claimed in claim 1, in which R³ is optionally substituted (C₆-C₁₄)-aryl, COOR⁴, R¹¹NH or CONHR⁴, where -NHR⁴ is the radical of an α -amino acid, its ω -amino-(C₂-C₈)-alkylamide, its (C₁-C₈)-alkyl ester, its (C₆-C₁₄)-aryl-(C₁-C₄)-alkyl ester or its derivative in which the carboxylic acid group is converted into the group Het-CO, in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

9. A compound of the formula I as claimed in claim 8, wherein the radical of the α -amino acids is selected from the group consisting of valine, lysine, phenylglycine, phenylalanine, tryptophan, their (C_1 - C_8)-alkyl esters, their (C_6 - C_{14})-aryl-(C_1 - C_4)-alkyl esters, and Het-CO derivatives; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.
10. A compound of the formula I as claimed in claimed 1, in which simultaneously
 W is R^1 -A-C(R^{13});
 Y is a carbonyl group;
 Z is $N(R^0)$;
 A is ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene, phenylenemethyl, methylenephenoxy or methylenephenoymethyl;
 B is an unsubstituted or substituted methylene radical;
 D is $C(R^2)(R^3)$;
 E is $R^{10}CO$;
 R is hydrogen or (C_1 - C_4)-alkyl;
 R^0 is (C_1 - C_8)-alkyl, (C_3 - C_8)-cycloalkyl, optionally substituted (C_6 - C_{14})-aryl or (C_6 - C_{14})-aryl-(C_1 - C_8)-alkyl optionally substituted in the aryl radical;
 R^1 is one of the radicals -O-C(O)- R^{21} , -O-C(O)-OR²², -O-C(O)-N(R^{21})- R^{28} , -N(R^{29})-C(O)-OR²², -N(R^{28})-C(O)-N(R^{21})- R^{28} , -N(R^{28})-C(S)-N(R^{21})- R^{28} or cyano;
 R^2 is hydrogen;
 R^3 is the radical CONHR⁴;
 R⁴ is methyl which is substituted by hydroxycarbonyl and a radical from the group consisting of (C_1 - C_4)-alkyl, phenyl and benzyl, or is methyl which is substituted by (C_1 - C_8)-alkoxycarbonyl and a radical from the group consisting of (C_1 - C_4)-alkyl, phenyl and benzyl, or is methyl which is substituted by Het-CO and a radical from the group consisting of (C_1 - C_4)-alkyl, phenyl and benzyl;
 R¹⁰ is hydroxyl or (C_1 - C_8)-alkoxy;
 R¹³ is (C_1 - C_6)-alkyl, (C_3 - C_7)-cycloalkyl or benzyl;
 b, c and d are 1 and e, f and g are 0;
 h is 1 or 2;
 in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.
35. 11. A compound of the formula I as claimed in claim 1, in which simultaneously
 W is R^1 -A-CH=C and therein A is a phenylene radical or a methylenephenoxy radical, or W is R^1 -A-C(R^{13}) and therein A is a bivalent radical from the group consisting of methylene, ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene, phenylenemethyl, methylenephenoxy, methylenephenoymethyl;
 B is a bivalent radical from the group consisting of methylene, ethylene, trimethylene, tetramethylene, vinylene, phenylene or substituted methylene or ethylene;
 E is $R^{10}CO$;

R is hydrogen or (C₁-C₆)-alkyl;

R⁰ is (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical;

R¹ is one of the radicals -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸,

5 -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸,
-O-S(O)₂-N(R²¹)-R²⁸, -O-S(O)-N(R²¹)-R²⁸,

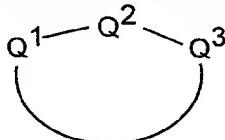
-N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(S)-R²¹,

-N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸,

-N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹,

10 -N(R²⁸)-S(O)₂-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-N(R²¹)-R²⁸,

-C(S)-R²¹, -C(S)-N(R²¹)-R²⁸ or cyano or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



in which

15 Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂;

Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-) representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

R² is hydrogen or (C₁-C₈)-alkyl;

R³ is CONHR¹⁵ or CONHR⁴ where R⁴ herein is a (C₁-C₈)-alkyl radical which is unsubstituted or substituted by one or more (C₆-C₁₄)-aryl radicals;

25 R¹⁵ is R¹⁶-(C₁-C₆)-alkyl or R¹⁶, where R¹⁶ is a 7- to 12-membered bridged bicyclic or tricyclic radical which is saturated or partially unsaturated and which can also contain one to four identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur and which can also be substituted by one or more identical or different substituents from the group consisting of (C₁-C₄)-alkyl and oxo;

30 and e, g and h independently of one another are the numbers 0, 1, 2 or 3 and b, c and d are 1; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

12. A compound of the formula I as claimed in claim 11, wherein R¹⁵ is an adamantyl radical or an adamantylmethyl radical; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

13. A compound of the formula I as claimed in claim 1, in which simultaneously

W is R¹-A-C(R¹³);

Y is a carbonyl group;

Z is N(R⁰);

A is ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene,
5 phenylenemethyl, methylenephenoxy or methylenephenoymethyl;

B is an unsubstituted or substituted methylene radical;

D is C(R²)(R³);

E is R¹⁰CO;

R is hydrogen or (C₁-C₄)-alkyl;

10 R⁰ is (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical;

R¹ is one of the radicals -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸,

-N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸ or cyano;

R² is hydrogen;

R³ is CONHR¹⁵ or CONHR⁴ where R⁴ herein is a (C₁-C₆)-alkyl radical which is unsubstituted or substituted by one or more (C₆-C₁₀)-aryl radicals;

R¹⁰ is hydroxyl or (C₁-C₈)-alkoxy;

R¹³ is (C₁-C₆)-alkyl, (C₃-C₈)-cycloalkyl or benzyl;

R¹⁵ is an adamantyl radical or an adamantylmethyl radical;

20 b, c and d are 1 and e, f and g are 0;

h is 1 or 2;

in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically
tolerable salts.

25 14. A compound of the formula I as claimed in claim 1, in which simultaneously

W is R¹-A-C(R¹³);

Y is a carbonyl group;

Z is N(R⁰);

A is ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene,
30 phenylenemethyl, methylenephenoxy or methylenephenoymethyl;

B is an unsubstituted or substituted methylene radical or ethylene radical;

D is C(R²)(R³);

E is R¹⁰CO;

R is hydrogen or (C₁-C₄)-alkyl;

35 R⁰ is (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl which is optionally substituted in the aryl radical;

R¹ is one of the radicals -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸,

-O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸,

-O-S(O)₂-N(R²¹)-R²⁸, -O-S(O)-N(R²¹)-R²⁸,

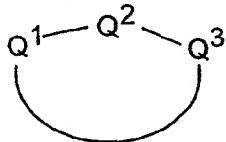
40 -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(S)-R²¹,

-N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸,

-N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹,

-N(R²⁸)-S(O)₂-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-N(R²¹)-R²⁸,

-C(S)-R²¹, -C(S)-N(R²¹)-R²⁸ or cyano or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



5 in which

Q¹ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂-;

Q³ is -C(R²¹)₂-, =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-)- representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

R² is hydrogen;

R³ is an unsubstituted phenyl radical or naphthyl radical, a phenyl radical or naphthyl radical substituted by one, two or three identical or different radicals from the group consisting of (C₁-C₄)-alkyl, (C₁-C₄)-alkoxy, hydroxyl, halogen, trifluoromethyl, nitro, methylenedioxy, ethylenedioxy, hydroxycarbonyl, (C₁-C₄)-alkoxycarbonyl, aminocarbonyl, cyano, phenyl, phenoxy and benzyloxy, a pyridyl radical, a (C₁-C₄)-alkyl radical, a (C₂-C₄)-alkenyl radical, a (C₂-C₄)-alkynyl radical or a (C₅-C₆)-cycloalkyl radical;

R¹⁰ is hydroxyl or (C₁-C₈)-alkoxy;

R¹³ is (C₁-C₆)-alkyl, (C₃-C₇)-cycloalkyl or benzyl;

b, c and d are 1 and e, f and g are 0;

h is 1 or 2;

in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

25 15. A compound of the formula I as claimed in claim 1, in which simultaneously

W is R¹-A-C(R¹³);

Y is a carbonyl group;

Z is N(R⁰);

A is ethylene, trimethylene, tetramethylene, pentamethylene, cyclohexylene, phenylene, phenylenemethyl, methylenephenoxy, methylenephenoymethyl;

B is an unsubstituted or substituted methylene radical or ethylene radical;

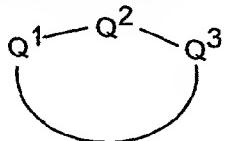
D is C(R²)(R³);

E is R¹⁰CO;

R is hydrogen or (C₁-C₄)-alkyl;

R⁰ is (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical;

R¹ is one of the radicals -S(O)-N(R²¹)-R²⁸, -S(O)₂-N(R²¹)-R²⁸, -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -O-C(S)-N(R²¹)-R²⁸, -O-S(O)₂-N(R²¹)-R²⁸, -O-S(O)-N(R²¹)-R²⁸, -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(S)-R²¹, -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸, -N(R²⁸)-S(O)₂-R²², -N(R²⁸)-S(O)-R²², -N(R²⁸)-S(O)₂-OR²¹, -N(R²⁸)-S(O)-OR²¹, -N(R²⁸)-S(O)-N(R²¹)-R²⁸, -N(R²⁸)-S(O)-N(R²¹)-R²⁸, -C(S)-R²¹, -C(S)-N(R²¹)-R²⁸ or cyano or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



in which

Q¹ is -C(R²¹)₂- =C(R²¹)-, -N(R²⁸)-, -O- or -S-;

Q² is -S(O)- or -S(O)₂;

Q³ is -C(R²¹)₂- =C(R²¹)-, -N(R²⁸)-, -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-)- representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

R² is hydrogen;

R³ is R¹¹NH;

R¹⁰ is hydroxyl or (C₁-C₈)-alkoxy;

R¹³ is (C₁-C₆)-alkyl, (C₃-C₇)-cycloalkyl or benzyl;

b, c, d and e are 1 and f and g are 0;

h is 0;

in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

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16. A compound of the formula I as claimed in claim 15, in which

R¹¹ is R^{12a}, R^{12a}-CO, R^{12a}-O-CO, R^{12b}-CO, R^{12b}-CS or R^{12a}-S(O)₂;

R^{12a} is (C₁-C₁₀)-alkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl

30 optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical or the radical R¹⁵;

R^{12b} is R^{12a}-NH;

in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

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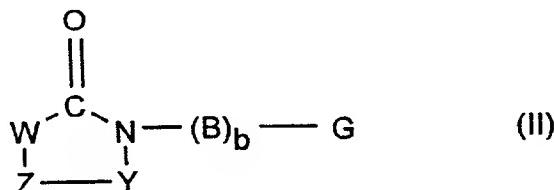
17. A compound of the formula I as claimed in claim 1, in which a substituted methylene radical or substituted ethylene radical representing the group B carries as a substituent a radical from the group consisting of (C₁-C₈)-alkyl, (C₂-C₆)-alkenyl, (C₂-C₆)-alkynyl, and

(C₃-C₈)-cycloalkyl, (C₃-C₈)-cycloalkyl-(C₁-C₄)-alkyl, optionally substituted (C₆-C₁₀)-aryl, (C₆-C₁₀)-aryl-(C₁-C₄)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, and heteroaryl-(C₁-C₄)-alkyl optionally substituted in the heteroaryl radical, in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

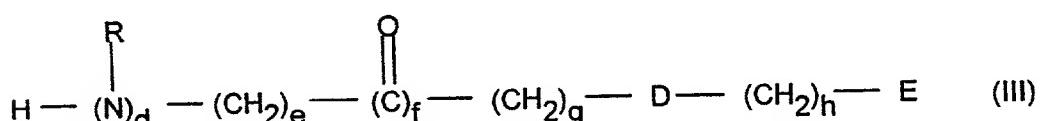
18. A compound of the formula I as claimed in claim 1, in which B is an unsubstituted methylene radical or a methylene radical which is substituted by a (C₁-C₈)-alkyl radical; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

19. A compound of the formula I as claimed in claim 1, in which R¹ is one of the radicals -O-C(O)-R²¹, -O-C(O)-OR²², -O-C(O)-N(R²¹)-R²⁸, -N(R²⁹)-C(O)-OR²², -N(R²⁸)-C(O)-N(R²¹)-R²⁸, -N(R²⁸)-C(S)-N(R²¹)-R²⁸ or cyano; in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts.

20. A process for the preparation of compounds of the formula I as claimed in claim 1, which comprises carrying out a fragment condensation of a compound of the formula II

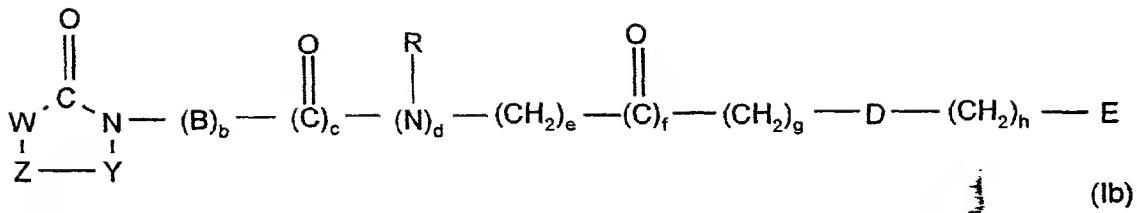


with a compound of the formula III



30 where W, Y, Z, B, D, E and R, and b, d, e, f, g, and h are defined as indicated in claim 1 and G is hydroxycarbonyl, (C₁-C₆)-alkoxycarbonyl, activated carboxylic acid derivatives.

21. A pharmaceutical preparation comprising an effective amount of one or more compounds of the formula Ib.



in which

W is $R^1-A-C(R^{13})$ or $R^1-A-CH=C$;

Y is a carbonyl, thiocarbonyl or methylene group;

Z is $N(R^0)$, oxygen, sulfur or a methylene group;

- A is a bivalent radical from the group consisting of (C_1-C_6)-alkylene, (C_3-C_{12})-cycloalkylene, (C_1-C_6)-alkylene-(C_3-C_{12})-cycloalkyl, phenylene, phenylene-(C_1-C_6)-alkyl, (C_1-C_6)-alkylenephenyl, (C_1-C_6)-alkylenephenyl-(C_1-C_6)-alkyl, phenylene-(C_2-C_6)-alkenyl or a bivalent radical of a 5- or 6-membered, saturated or unsaturated ring, which can contain 1 or 2 nitrogen atoms and can be mono- or disubstituted by (C_1-C_6)-alkyl or doubly bonded oxygen or sulfur or is a direct bond;
- B is a bivalent radical from the group consisting of (C_1-C_6)-alkylene, (C_2-C_6)-alkenylene, phenylene, phenylene-(C_1-C_3)-alkyl, (C_1-C_3)-alkylenephenyl where the bivalent (C_1-C_6)-alkylene radical can be unsubstituted or substituted by a radical from the group consisting of (C_1-C_8)-alkyl, (C_2-C_8)-alkenyl, (C_2-C_8)-alkynyl, (C_3-C_{10})-cycloalkyl, (C_3-C_{10})-cycloalkyl-(C_1-C_6)-alkyl, optionally substituted (C_6-C_{14})-aryl, (C_6-C_{14})-aryl-(C_1-C_6)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl and heteroaryl-(C_1-C_6)-alkyl optionally substituted in the heteroaryl radical;
- D is $C(R^2)(R^3)$, $N(R^3)$ or $CH=C(R^3)$;
- E is tetrazolyl, $(R^8O)_2P(O)$, $HOS(O)_2$, $R^9NHS(O)_2$ or $R^{10}CO$;
- R is hydrogen, (C_1-C_8)-alkyl, (C_3-C_{12})-cycloalkyl, (C_3-C_{12})-cycloalkyl-(C_1-C_8)-alkyl, optionally substituted (C_6-C_{14})-aryl, (C_6-C_{14})-aryl-(C_1-C_8)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C_1-C_8)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be mono- or polysubstituted by fluorine;
- R^0 is hydrogen, (C_1-C_8)-alkyl, (C_3-C_{12})-cycloalkyl, (C_3-C_{12})-cycloalkyl-(C_1-C_8)-alkyl, (C_6-C_{12})-bicycloalkyl, (C_6-C_{12})-bicycloalkyl-(C_1-C_8)-alkyl, (C_6-C_{12})-tricycloalkyl, (C_6-C_{12})-tricycloalkyl-(C_1-C_8)-alkyl, optionally substituted (C_6-C_{14})-aryl, (C_6-C_{14})-aryl-(C_1-C_8)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-(C_1-C_8)-alkyl optionally substituted in the heteroaryl radical, CHO , (C_1-C_8)-alkyl-CO, (C_3-C_{12})-cycloalkyl-CO, (C_3-C_{12})-cycloalkyl-(C_1-C_8)-alkyl-CO, (C_6-C_{12})-bicycloalkyl-CO, (C_6-C_{12})-bicycloalkyl-(C_1-C_8)-alkyl-CO, (C_6-C_{12})-tricycloalkyl-CO, (C_6-C_{12})-tricycloalkyl-(C_1-C_8)-alkyl-CO, optionally substituted (C_6-C_{14})-aryl-CO, (C_6-C_{14})-aryl-(C_1-C_8)-alkyl-CO optionally substituted in the aryl radical, optionally substituted heteroaryl-CO, heteroaryl-(C_1-C_8)-alkyl-CO optionally substituted in the heteroaryl radical, (C_1-C_8)-alkyl-S(O)_n, (C_3-C_{12})-cycloalkyl-S(O)_n, (C_3-C_{12})-cycloalkyl-(C_1-C_8)-alkyl-S(O)_n, (C_6-C_{12})-bicycloalkyl-S(O)_n, (C_6-C_{12})-bicycloalkyl-(C_1-C_8)-alkyl-

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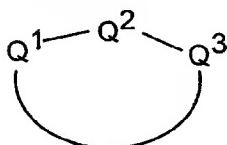
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$S(O)_n$, (C_6-C_{12}) -tricycloalkyl- $S(O)_n$, (C_6-C_{12}) -tricycloalkyl-(C_1-C_8)-alkyl- $S(O)_n$, optionally substituted (C_6-C_{14}) -aryl- $S(O)_n$, (C_6-C_{14}) -aryl-(C_1-C_8)-alkyl- $S(O)_n$ optionally substituted in the aryl radical, optionally substituted heteroaryl- $S(O)_n$ or heteroaryl-(C_1-C_8)-alkyl- $S(O)_n$ optionally substituted in the heteroaryl radical, where n is 1 or 2;

- 5 R^1 is one of the radicals $-S-R^{21}$, $-S-S-R^{21}$, $-S(O)-R^{22}$, $-S(O)_2-R^{22}$,
 $-S-OR^{21}$, $-S(O)-OR^{21}$, $-S(O)_2-OR^{21}$, $-S-N(R^{21})-R^{28}$,
 $-S(O)-N(R^{21})-R^{28}$, $-S(O)_2-N(R^{21})-R^{28}$, $-S-C(O)-R^{21}$,
 $-S-C(O)-OR^{22}$, $-S-C(S)-SR^{22}$, $-S-C(O)-N(R^{21})-R^{28}$,
 $-S-C(S)-N(R^{21})-R^{28}$,
- 10 $-O-C(O)-R^{21}$, $-O-C(S)-R^{21}$, $-O-C(O)-OR^{22}$, $-O-C(O)-N(R^{21})-R^{28}$,
 $-O-C(S)-N(R^{21})-R^{28}$, $-O-S(O)_2-OR^{21}$, $-O-S(O)-OR^{21}$, $-O-S(O)_2-N(R^{21})-R^{28}$,
 $-O-S(O)-N(R^{21})-R^{28}$, $-O-S(O)_2-R^{22}$, $-O-S(O)-R^{22}$,
 $-O-P(O)(OR^{21})_2$, $-O-P(O)(OR^{21})-N(R^{21})-R^{28}$, $-O-P(O)(N(R^{21})-R^{28})_2$,
 $-N(R^{29})-C(O)-OR^{22}$, $-N(R^{28})-C(O)-SR^{22}$, $-N(R^{28})-C(S)-OR^{22}$,
 $-N(R^{28})-C(S)-SR^{22}$, $-N(R^{28})-C(S)-R^{21}$,
 $-N(R^{28})-C(O)-N(R^{21})-R^{28}$, $-N(R^{28})-C(S)-N(R^{21})-R^{28}$,
 $-N(R^{28})-S(O)_2-R^{22}$, $-N(R^{28})-S(O)-R^{22}$, $-N(R^{28})-S(O)_2-OR^{21}$, $-N(R^{28})-S(O)-OR^{21}$,
 $-N(R^{28})-S(O)_2-N(R^{21})-R^{28}$, $-N(R^{28})-S(O)-N(R^{21})-R^{28}$,
 $-N(R^{28})-P(O)(OR^{21})_2$, $-N(R^{28})-P(O)(OR^{21})-N(R^{21})-R^{28}$,
 $-N(R^{28})-P(O)(N(R^{21})-R^{28})_2$, $-N(R^{28})-P(O)(R^{22})-OR^{21}$,
 $-N(R^{28})-P(O)(R^{22})-N(R^{21})-R^{28}$, $-N(R^{28})-P(O)(R^{22})_2$,
 $-P(O)(OR^{21})_2$, $-P(O)(OR^{21})-N(R^{21})-R^{28}$, $-P(O)(N(R^{21})-R^{28})_2$,
 $-P(O)(R^{22})-OR^{21}$,
 $-P(O)(R^{22})-N(R^{21})-R^{28}$, $-P(O)(R^{22})_2$,
 $-C(S)-R^{21}$, $-C(S)-SR^{21}$, $-C(S)-N(R^{21})-R^{28}$,
- 25 cyano, halogen, nitro or methylenedioxy or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula



30 in which

Q^1 is $-C(R^{21})_2-$, $=C(R^{21})-$, $-N(R^{28})-$, $-O-$ or $-S-$;

Q^2 is $-S(O)-$ or $-S(O)_2-$;

Q^3 is $-C(R^{21})_2-$, $=C(R^{21})-$, $-N(R^{28})-$, $-O-$, $-S-$, $-C(R^{21})(-)-$ or $-N(-)-$,

35 where the heterocyclic ring can be bonded to the group A via the free bond in the groups $-C(R^{21})(-)-$ or $-N(-)-$ representing Q^3 or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

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- 5 R² is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl;
- R³ is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, (C₃-C₈)-cycloalkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₂-C₈)-alkenylcarbonyl, (C₂-C₈)-alkynylcarbonyl, pyridyl, R¹¹NH, R⁴CO, COOR⁴, CON(CH₃)R⁴, CONHR⁴, CSNHR⁴, COOR¹⁵, CON(CH₃)R¹⁵ or CONHR¹⁵;
- 10 R⁴ is hydrogen or (C₁-C₂₈)-alkyl which can optionally be mono- or polysubstituted by identical or different radicals from the group consisting of hydroxyl, hydroxycarbonyl, aminocarbonyl, mono- or di-((C₁-C₁₈)-alkyl)aminocarbonyl, amino-(C₂-C₁₈)-alkylaminocarbonyl, amino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₂-C₁₈)-alkylaminocarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxycarbonyl which can also be substituted in the aryl radical, amino, mercapto, (C₁-C₁₈)-alkoxy, (C₁-C₁₈)-alkoxycarbonyl, Het-CO, optionally substituted (C₃-C₈)-cycloalkyl, HOS(O)₂-(C₁-C₃)-alkyl, R⁹NHS(O)₂-(C₁-C₃)-alkyl, (R⁸O)₂P(O)-(C₁-C₃)-alkyl, tetrazolyl-(C₁-C₃)-alkyl, halogen, nitro, trifluoromethyl or the radical R⁵;
- R⁵ is optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, a mono- or bicyclic 5- to 12-membered heterocyclic ring which can be aromatic, partially hydrogenated or completely hydrogenated and which can contain one, two or three identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur, a radical R⁶ or a radical R⁶CO-, where the aryl radical and, independently thereof, the heterocyclic radical can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₁₈)-alkyl, (C₁-C₁₈)-alkoxy, halogen, nitro, amino and trifluoromethyl;
- 30 R⁶ is R'R⁸N, R⁷O or R⁷S or an amino acid side chain, a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated)azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or in which the peptide bond can be reduced to -NH-CH₂-, and its esters and amides, where hydrogen or hydroxymethyl can optionally stand in place of free functional groups and/or where free functional groups can be protected by protective groups customary in peptide chemistry;
- 35 R⁷ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted, preferably monosubstituted, by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated)azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or in which the peptide bond can be reduced to -NH-CH₂-;
- 40

	R ⁸	is hydrogen, (C ₁ -C ₁₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl or (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl which can also be substituted in the aryl radical;
5	R ⁹	is hydrogen, aminocarbonyl, (C ₁ -C ₁₈)-alkylaminocarbonyl, (C ₃ -C ₈)-cycloalkylaminocarbonyl, optionally substituted (C ₆ -C ₁₄)-arylamino carbonyl, (C ₁ -C ₁₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl or (C ₃ -C ₈)-cycloalkyl;
10	R ¹⁰	is hydroxyl, (C ₁ -C ₁₈)-alkoxy, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkoxy which can also be substituted in the aryl radical, optionally substituted (C ₆ -C ₁₄)-aryloxy, amino or mono- or di-((C ₁ -C ₁₈)-alkyl)amino;
15	R ¹¹	is hydrogen, R ^{12a} , R ^{12a} -CO, H-CO, R ^{12a} -O-CO, R ^{12b} -CO, R ^{12b} -CS, R ^{12a} -S(O) ₂ or R ^{12b} -S(O) ₂ ;
20	R ^{12a}	is (C ₁ -C ₁₈)-alkyl, (C ₂ -C ₈)-alkenyl, (C ₂ -C ₈)-alkynyl, (C ₃ -C ₁₂)-cycloalkyl, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-(C ₁ -C ₈)-alkyl optionally substituted in the heteroaryl radical or the radical R ¹⁵ ;
25	R ^{12b}	is amino, di-((C ₁ -C ₁₈)-alkyl)-amino or R ^{12a} -NH;
30	R ¹³	is hydrogen, (C ₁ -C ₆)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, (C ₃ -C ₈)-cycloalkyl or (C ₃ -C ₈)-cycloalkyl-(C ₁ -C ₈)-alkyl;
35	R ¹⁵	is R ¹⁶ -(C ₁ -C ₆)-alkyl or R ¹⁶ ;
40	R ¹⁶	is a 6- to 24-membered, bicyclic or tricyclic radical which is saturated or partially unsaturated and which can also contain one, two, three or four identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur and which can also be substituted by one or more identical or different substituents from the group consisting of (C ₁ -C ₄)-alkyl and oxo;
	R ²¹	is hydrogen, (C ₁ -C ₈)-alkyl, hydroxy-(C ₁ -C ₈)-alkyl, (C ₂ -C ₈)-alkenyl, (C ₃ -C ₁₂)-cycloalkyl, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C ₁ -C ₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can also be monosubstituted or polysubstituted by fluorine and the radicals R ²¹ can be identical or different if they occur two or more times;
	R ²²	is (C ₁ -C ₈)-alkyl, hydroxy-(C ₁ -C ₈)-alkyl, (C ₂ -C ₈)-alkenyl, (C ₃ -C ₁₂)-cycloalkyl, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C ₁ -C ₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be monosubstituted or polysubstituted by fluorine and the radicals R ²² can be identical or different if they occur two or more times;
	R ²⁸	is one of the radicals R ²¹ -, R ²¹ N(R ²¹)-, R ²¹ C(O)-, R ²² O-C(O)-, R ²¹ N(R ²¹)-C(O)- or R ²¹ N(R ²¹)-C(=N(R ²¹))-
	R ²⁹	is one of the radicals R ²² -, R ²¹ N(R ²¹)-, R ²¹ C(O)-, R ²² O-C(O)-, R ²¹ N(R ²¹)-C(O)- or R ²¹ N(R ²¹)-C(=N(R ²¹))-
	Het	is the radical of a 5- to 10-membered, monocyclic or polycyclic heterocycle bonded via

a nitrogen atom, which can be aromatic or partially unsaturated or saturated and which can contain one, two, three or four identical or different additional ring heteroatoms from the group consisting of oxygen, nitrogen and sulfur and which can be optionally substituted on carbon atoms and on additional ring nitrogen atoms, where there can be identical or different radicals R^h, R^hCO or R^hO-CO as substituents on additional ring nitrogen atoms and R^h is hydrogen, (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, (C₃-C₈)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical;

b, c, d and f independently of one another can be 0 or 1, but cannot all simultaneously be 0;
 e, g and h independently of one another are 0, 1, 2, 3, 4, 5 or 6;
 in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts; and one or more pharmaceutically innocuous carriers and/or additives.

22. A pharmaceutical preparation as claimed in claim 21 wherein the effective amount is an amount effective for the suppression of inflammation.

23. A pharmaceutical preparation as claimed in claim 21 wherein the effective amount is an amount effective for the suppression of leucocyte migration and/or adhesion.

24. A pharmaceutical preparation as claimed in claim 21 wherein the effective amount is an amount effective for treatment or prevention of a disease selected from the group consisting of rheumatoid arthritis, inflammatory bowel disease, systemic lupus erythematosus, inflammatory disorders of the central nervous system, asthma, allergies, cardiovascular disorders, arteriosclerosis, restenoses, diabetes, damage to organ transplants, tumor growth tumor metastasis, and malaria.

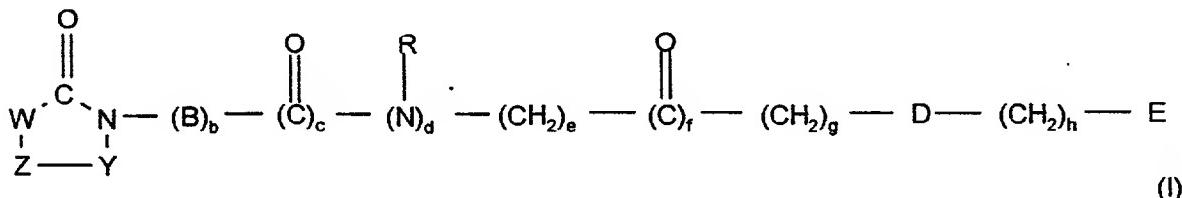
25. A method for the suppression of inflammation comprising administering to a subject in need thereof an effective amount of the preparation as claimed in claim 21.

26. A method for antagonizing VLA-4 comprising administering to a subject in need thereof an effective amount of the preparation as claimed in claim 21.

27. A method for inhibiting leucocyte adhesion and/or leucocyte migration comprising administering to a subject in need thereof an effective amount of the preparation as claimed in claim 21.

28. A method for treatment or prevention of a disease or disorder selected from the group consisting of rheumatoid arthritis, inflammatory bowel disease, systemic lupus erythematosus, inflammatory disorders of the central nervous system, asthma, allergies, cardiovascular disorders, arteriosclerosis, restenoses, diabetes, damage to organ transplants, tumor growth, tumor metastasis, and malaria comprising administering to a subject in need thereof an effective amount of the preparation as claimed in claim 21.

29. The preparation of claim 21 wherein the effective amount is a VLA-4-antagonizing effective amount.
 30. The pharmaceutical preparation as claimed in claim 29 wherein the VLA-4-antagonizing effective amount is an amount effective for the suppression of inflammation.
 31. The pharmaceutical preparation as claimed in claim 29 wherein the VLA-4-antagonizing effective amount is an amount effective for suppression of leucocyte adhesion and/or migration.
 32. The pharmaceutical preparation as claimed in claim 29 wherein the VLA-4-antagonizing effective amount is an amount effective for treating or preventing a disease or disorder selected from the group consisting of rheumatoid arthritis, inflammatory bowel disease, systemic lupus erythematosus, inflammatory disorders of the central nervous system, asthma, allergies, cardiovascular disorders, arteriosclerosis, restenoses, diabetes, damage to organ transplants, tumor growth, tumor metastasis, and malaria.
 33. A kit comprising a VLA-4-antagonizing effective amount of one or more compounds of the formula 1b



in which

W is $R^1-A-C(R^{13})$ or $R^1-A-CH=C$:

Y is a carbonyl, thiocarbonyl or methylene group;

Z is N(R^0), oxygen, sulfur or a methylene group;

A is a bivalent radical from the group consisting of (C_1 - C_6)-alkylene, (C_3 - C_{12})-cycloalkylene, (C_1 - C_6)-alkylene-(C_3 - C_{12})-cycloalkyl, phenylene, phenylene-(C_1 - C_6)-alkyl, (C_1 - C_6)-alkylenephenyl, (C_1 - C_6)-alkylenephenyl-(C_1 - C_6)-alkyl, phenylene-(C_2 - C_6)-alkenyl or a bivalent radical of a 5- or 6-membered saturated or unsaturated ring which can contain 1 or 2 nitrogen atoms and can be mono- or disubstituted by (C_1 - C_6)-alkyl or doubly bonded oxygen or sulfur, or is a direct bond;

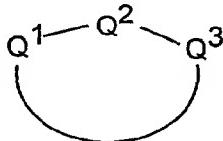
B is a bivalent radical from the group consisting of (C_1 - C_6)-alkylene, (C_2 - C_6)-alkenylene, phenylene, phenylene- $(C_1$ - C_3)-alkyl, (C_1 - C_3)-alkylenephenoxy, where the bivalent (C_1 - C_6)-alkylene radical can be unsubstituted or substituted by a radical from the group consisting of (C_1 - C_8)-alkyl, (C_2 - C_8)-alkenyl, (C_2 - C_8)-alkynyl, (C_3 - C_{10})-cycloalkyl, (C_3 - C_{10})-cycloalkyl-(C_1 - C_6)-alkyl, optionally substituted (C_6 - C_{14})-aryl, (C_6 - C_{14})-aryl-

		(C ₁ -C ₆)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl and heteroaryl-(C ₁ -C ₆)-alkyl optionally substituted in the heteroaryl radical;
D		is C(R ²)(R ³), N(R ³) or CH=C(R ³);
E		is tetrazolyl, (R ⁸ O) ₂ P(O), HOS(O) ₂ , R ⁹ NHS(O) ₂ or R ¹⁰ CO;
5	R	is hydrogen, (C ₁ -C ₈)-alkyl, (C ₃ -C ₁₂)-cycloalkyl, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl and heteroaryl-(C ₁ -C ₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be mono- or polysubstituted by fluorine;
10	R ⁰	is hydrogen, (C ₁ -C ₈)-alkyl, (C ₃ -C ₁₂)-cycloalkyl, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl, (C ₆ -C ₁₂)-bicycloalkyl, (C ₆ -C ₁₂)-bicycloalkyl-(C ₁ -C ₈)-alkyl, (C ₆ -C ₁₂)-tricycloalkyl, (C ₆ -C ₁₂)-tricycloalkyl-(C ₁ -C ₈)-alkyl, optionally substituted (C ₆ -C ₁₄)-aryl, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-(C ₁ -C ₈)-alkyl optionally substituted in the heteroaryl radical, CHO, (C ₁ -C ₈)-alkyl-CO, (C ₃ -C ₁₂)-cycloalkyl-CO, (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl-CO, (C ₆ -C ₁₂)-bicycloalkyl-CO, (C ₆ -C ₁₂)-bicycloalkyl-(C ₁ -C ₈)-alkyl-CO, (C ₆ -C ₁₂)-tricycloalkyl-CO, (C ₆ -C ₁₂)-tricycloalkyl-(C ₁ -C ₈)-alkyl-CO, optionally substituted (C ₆ -C ₁₄)-aryl-CO, (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl-CO optionally substituted in the aryl radical, optionally substituted heteroaryl-CO, heteroaryl-(C ₁ -C ₈)-alkyl-CO optionally substituted in the heteroaryl radical, (C ₁ -C ₈)-alkyl-S(O) _n , (C ₃ -C ₁₂)-cycloalkyl-S(O) _n , (C ₃ -C ₁₂)-cycloalkyl-(C ₁ -C ₈)-alkyl-S(O) _n , (C ₆ -C ₁₂)-bicycloalkyl-1-S(O) _n , (C ₆ -C ₁₂)-bicycloalkyl-(C ₁ -C ₈)-alkyl-S(O) _n , (C ₆ -C ₁₂)-tricycloalkyl-S(O) _n , (C ₆ -C ₁₂)-tricycloalkyl-(C ₁ -C ₈)-alkyl-S(O) _n , optionally substituted (C ₆ -C ₁₄)-aryl-S(O) _n , (C ₆ -C ₁₄)-aryl-(C ₁ -C ₈)-alkyl-S(O) _n optionally substituted in the aryl radical, optionally substituted heteroaryl-S(O) _n or heteroaryl-(C ₁ -C ₈)-alkyl-S(O) _n optionally substituted in the heteroaryl radical, where n is 1 or 2;
15	R ¹	is one of the radicals -S-R ²¹ , -S-S-R ²¹ , -S(O)-R ²² , -S(O) ₂ -R ²² , -S-OR ²¹ , -S(O)-OR ²¹ , -S(O) ₂ -OR ²¹ , -S-N(R ²¹)-R ²⁸ , -S(O)-N(R ²¹)-R ²⁸ , -S(O) ₂ -N(R ²¹)-R ²⁸ , -S-C(O)-R ²¹ , -S-C(O)-OR ²² , -S-C(S)-SR ²² , -S-C(O)-N(R ²¹)-R ²⁸ , -S-C(S)-N(R ²¹)-R ²⁸ , -O-C(O)-R ²¹ , -O-C(S)-R ²¹ , -O-C(O)-OR ²² , -O-C(O)-N(R ²¹)-R ²⁸ , -O-C(S)-N(R ²¹)-R ²⁸ , -O-S(O) ₂ -OR ²¹ , -O-S(O)-N(R ²¹)-R ²⁸ , -O-S(O) ₂ -R ²² , -O-S(O)-R ²² , -O-P(O)(OR ²¹) ₂ , -O-P(O)(OR ²¹)-N(R ²¹)-R ²⁸ , -O-P(O)(N(R ²¹)-R ²⁸) ₂ , -N(R ²⁹)-C(O)-OR ²² , -N(R ²⁸)-C(O)-SR ²² , -N(R ²⁸)-C(S)-OR ²² , -N(R ²⁸)-C(S)-SR ²² , -N(R ²⁸)-C(S)-R ²¹ , -N(R ²⁸)-C(O)-N(R ²¹)-R ²⁸ , -N(R ²⁸)-C(S)-N(R ²¹)-R ²⁸ , -N(R ²⁸)-S(O) ₂ -R ²² , -N(R ²⁸)-S(O)-R ²² , -N(R ²⁸)-S(O) ₂ -OR ²¹ , -N(R ²⁸)-S(O)-OR ²¹ , -N(R ²⁸)-S(O) ₂ -N(R ²¹)-R ²⁸ , -N(R ²⁸)-S(O)-N(R ²¹)-R ²⁸ , -N(R ²⁸)-P(O)(OR ²¹) ₂ , -N(R ²⁸)-P(O)(OR ²¹)-N(R ²¹)-R ²⁸ , -N(R ²⁸)-P(O)(N(R ²¹)-R ²⁸) ₂ , -N(R ²⁸)-P(O)(R ²²)-OR ²¹ , -N(R ²⁸)-P(O)(R ²²)-N(R ²¹)-R ²⁸ , -N(R ²⁸)-P(O)(R ²²) ₂ , -P(O)(OR ²¹) ₂ , -P(O)(OR ²¹)-N(R ²¹)-R ²⁸ , -P(O)(N(R ²¹)-R ²⁸) ₂ , -P(O)(R ²²)-OR ²¹ , -P(O)(R ²²)-N(R ²¹)-R ²⁸ , -P(O)(R ²²) ₂ ,
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-C(S)-R²¹, -C(S)-SR²¹, -C(S)-N(R²¹)-R²⁸,

cyano, halogen, nitro or methylenedioxy or the radical of an optionally substituted, 5- to 14-membered, mono- or polycyclic, saturated or unsaturated heterocyclic ring of the formula

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in which

Q¹ is -C(R²¹)₂-; =C(R²¹)-; -N(R²⁸)-; -O- or -S-;

Q² is -S(O)- or -S(O)₂-;

Q³ is -C(R²¹)₂-; =C(R²¹)-; -N(R²⁸)-; -O-, -S-, -C(R²¹)(-) or -N(-)-,

where the heterocyclic ring can be bonded to the group A via the free bond in the groups -C(R²¹)(-) or -N(-) representing Q³ or via any other desired ring carbon atom and where, if the heterocyclic ring is bonded to a ring system contained in the group A, the heterocyclic ring can also be fused via two adjacent atoms to the ring system in the group A;

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R² is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical or (C₃-C₈)-cycloalkyl;

R³ is hydrogen, (C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, (C₃-C₈)-cycloalkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₂-C₈)-alkenylcarbonyl, (C₂-C₈)-alkynylcarbonyl, pyridyl, R¹¹NH, R⁴CO, COOR⁴, CON(CH₃)R⁴, CONHR⁴, CSNHR⁴, COOR¹⁵, CON(CH₃)R¹⁵ or CONHR¹⁵;

R⁴ is hydrogen or (C₁-C₂₈)-alkyl which can optionally be mono- or polysubstituted by identical or different radicals from the group consisting of hydroxyl, hydroxycarbonyl, aminocarbonyl, mono- or di-((C₁-C₁₈)-alkyl)aminocarbonyl, amino-(C₂-C₁₈)-alkylaminocarbonyl, amino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl,

(C₁-C₁₈)-alkylcarbonylamino-(C₁-C₃)-alkylphenyl-(C₁-C₃)-alkylaminocarbonyl, (C₁-C₁₈)-alkylcarbonylamino-(C₂-C₁₈)-alkylaminocarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxycarbonyl which can also be substituted in the aryl radical, amino, mercapto, (C₁-C₁₈)-alkoxy, (C₁-C₁₈)-alkoxycarbonyl, Het-CO, optionally substituted

(C₃-C₈)-cycloalkyl, HOS(O)₂-(C₁-C₃)-alkyl, R⁹NHS(O)₂-(C₁-C₃)-alkyl, (R⁸O)₂P(O)-

(C₁-C₃)-alkyl, tetrazolyl-(C₁-C₃)-alkyl, halogen, nitro, trifluoromethyl or the radical R⁵;

R⁵ is optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, a mono- or bicyclic 5- to 12-membered heterocyclic ring which can be aromatic, partially hydrogenated or completely hydrogenated and which can contain one, two or three identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur, a radical R⁶ or a radical R⁶CO-, where the aryl radical and, independently thereof, the heterocyclic radical can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₁₈)-alkyl, (C₁-C₁₈)-alkoxy, halogen, nitro, amino and trifluoromethyl;

- DETAILED DESCRIPTION
- 5 R⁶ is R⁷R⁸N, R⁷O or R⁷S or an amino acid side chain, a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-, and their esters and amides, where hydrogen or hydroxymethyl can optionally stand in place of free functional groups and/or where free functional groups can be protected by protective groups customary in peptide chemistry;
- 10 R⁷ is hydrogen, (C₁-C₁₈)-alkyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl, (C₁-C₁₈)-alkylcarbonyl, (C₁-C₁₈)-alkoxycarbonyl, (C₆-C₁₄)-arylcarbonyl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkylcarbonyl or (C₆-C₁₄)-aryl-(C₁-C₁₈)-alkyloxycarbonyl, where the alkyl groups can optionally be substituted by an amino group and/or where the aryl radicals can be mono- or polysubstituted by identical or different radicals from the group consisting of (C₁-C₈)-alkyl, (C₁-C₈)-alkoxy, halogen, nitro, amino and trifluoromethyl, or is a natural or unnatural amino acid, imino acid, optionally N-(C₁-C₈)-alkylated or N-((C₆-C₁₄)-aryl-(C₁-C₈)-alkylated) azaamino acid or a dipeptide radical which can also be substituted in the aryl radical and/or reduced in the peptide bond to -NH-CH₂-;
- 15 R⁸ is hydrogen, (C₁-C₁₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl which can also be substituted in the aryl radical;
- 20 R⁹ is hydrogen, aminocarbonyl, (C₁-C₁₈)-alkylaminocarbonyl, (C₃-C₈)-cycloalkylaminocarbonyl, optionally substituted (C₆-C₁₄)-arylaminocarbonyl, (C₁-C₁₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₃-C₈)-cycloalkyl;
- 25 R¹⁰ is hydroxyl, (C₁-C₁₈)-alkoxy, (C₆-C₁₄)-aryl-(C₁-C₈)-alkoxy which can also be substituted in the aryl radical, optionally substituted (C₆-C₁₄)-aryloxy, amino or mono- or di-((C₁-C₁₈)-alkyl)amino;
- 30 R¹¹ is hydrogen, R^{12a}, R^{12a}-CO, H-CO, R^{12a}-O-CO, R^{12b}-CO, R^{12b}-CS, R^{12a}-S(O)₂ or R^{12b}-S(O);
- 35 R^{12a} is (C₁-C₁₈)-alkyl, (C₂-C₈)-alkenyl, (C₂-C₈)-alkynyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl, heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical or the radical R¹⁵;
- 40 R^{12b} is amino, di-((C₁-C₁₈)-alkyl)amino or R^{12a}-NH;
- 45 R¹³ is hydrogen, (C₁-C₆)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, (C₃-C₈)-cycloalkyl or (C₃-C₈)-cycloalkyl-(C₁-C₈)-alkyl;
- 50 R¹⁵ is R¹⁶-(C₁-C₆)-alkyl or R¹⁶;
- 55 R¹⁶ is a 6- to 24-membered bicyclic or tricyclic radical which is saturated or partially unsaturated and which can also contain one, two, three or four identical or different heteroatoms from the group consisting of nitrogen, oxygen and sulfur and which can also be substituted by one or more identical or different substituents from the group consisting of (C₁-C₄)-alkyl and oxo;
- 60 R²¹ is hydrogen, (C₁-C₈)-alkyl, hydroxy-(C₁-C₈)-alkyl, (C₂-C₈)-alkenyl, (C₃-C₁₂)-cycloalkyl,

(C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can also be monosubstituted or polysubstituted by fluorine and the radicals R²¹

5 can be identical or different if they occur two or more times;

R²² is (C₁-C₈)-alkyl, hydroxy-(C₁-C₈)-alkyl, (C₂-C₈)-alkenyl, (C₃-C₁₂)-cycloalkyl, (C₃-C₁₂)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl, (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical, optionally substituted heteroaryl or heteroaryl-(C₁-C₈)-alkyl optionally substituted in the heteroaryl radical, where alkyl radicals can be monosubstituted or polysubstituted by fluorine and the radicals R²² can

10 be identical or different if they occur two or more times;

R²⁸ is one of the radicals R²¹-, R²¹N(R²¹)-, R²¹C(O)-, R²²O-C(O)-, R²¹N(R²¹)-C(O)- or R²¹N(R²¹)-C(=N(R²¹))-;

R²⁹ is one of the radicals R²²-, R²¹N(R²¹)-, R²¹C(O)-, R²²O-C(O)-, R²¹N(R²¹)-C(O)- or R²¹N(R²¹)-C(=N(R²¹))-;

Het is the radical of a 5- to 10-membered, monocyclic or polycyclic heterocycle bonded via a nitrogen atom, which can be aromatic or partially unsaturated or saturated and which can contain one, two, three or four identical or different additional ring heteroatoms from the group consisting of oxygen, nitrogen and sulfur and which can be optionally substituted on carbon atoms and on additional ring nitrogen atoms, where there can be identical or different radicals R^h, R^hCO or R^hO-CO as substituents on additional ring nitrogen atoms and R^h is hydrogen, (C₁-C₈)-alkyl, (C₃-C₈)-cycloalkyl, (C₃-C₈)-cycloalkyl-(C₁-C₈)-alkyl, optionally substituted (C₆-C₁₄)-aryl or (C₆-C₁₄)-aryl-(C₁-C₈)-alkyl optionally substituted in the aryl radical;

25 b, c, d and f independently of one another are 0 or 1, but cannot all simultaneously be 0;

e, g and h independently of one another are 0, 1, 2, 3, 4, 5 or 6;

in all its stereoisomeric forms and mixtures thereof in any ratio, and/or its physiologically tolerable salts;

30 instructions for use; and
one or more pharmaceutically innocuous carriers and/or additives.

34. The kit as claimed in claim 33 wherein the VLA-4-antagonizing effective amount is an amount effective for the suppression of inflammation.

35. The kit as claimed in claim 33 wherein the VLA-4-antagonizing effective amount is an amount effective for suppression of leucocyte adhesion and/or migration.

40 36. The kit as claimed in claim 33 wherein the VLA-4-antagonizing effective amount is an amount effective for treating or preventing a disease or disorder selected from the group consisting of rheumatoid arthritis, inflammatory bowel disease, systemic lupus erythematosus, inflammatory disorders of the central nervous system, asthma, allergies,

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cardiovascular disorders, arteriosclerosis, restenoses, diabetes, damage to organ transplants, tumor growth, tumor metastasis, and malaria.